

Inventor: Leonard KATZ and Peter REVILL Title: PRODUCTION OF POLYKETIDES

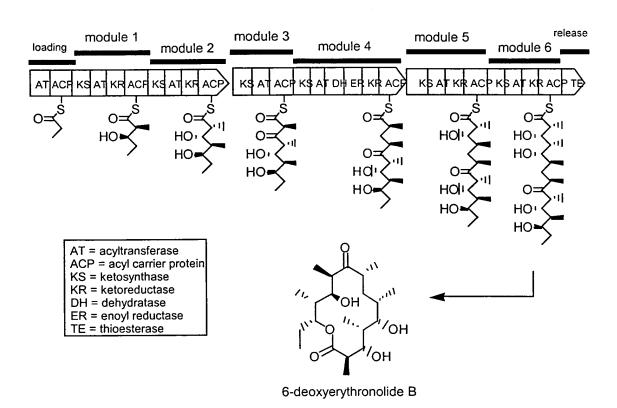


Figure 1



App No.: 10/607,809

Docket No.: 300622004810

Inventor: Leonard KATZ and Peter REVILL Title: PRODUCTION OF POLYKETIDES

Figure 2



App No.: 10/607,809 Docke Inventor: Leonard KATZ and Peter REVILL Title: PRODUCTION OF POLYKETIDES

gaac gaaa tcat	tcct ccgc gtag	ta to cc ag cc ag	ccgg gcgg gatt	acgae agcge tcgge	c tag c ag c at	gacci agat aacti tcgg	tggt cctc ggtc ccta	acc cac	ctaa tgat gatc qcqt	gcg cag ccg gct	gaga gtaa attc ccgc gc g	ggato gtgao ggtgo tg g	ct t cc a at c	catto tgcg	atgtg cgcgc cgatg tgtcc gttag ac sp	180 240
gtg Val 5	gaa Glu	ctc Leu	gcg Ala	gac Asp	agg Arg 10	gct Ala	cga Arg	cga Arg	cgc Arg	gcg Ala 15	tgc Cys	cgg Arg	ctg Leu	ctc Leu	agg Arg 20	402
cgt Arg	tgg Trp	ctg Leu	acc	aI gag Glu 25	acg Thr	cac His	act Thr	ccg Pro	gtg Val 30	gag Glu	ccc Pro	ggc Gly	ccg Pro	ctg Leu 35	tcc Ser	450
ctg Leu	cgg Arg	atc Ile	ggc Gly 40	ccg Pro	gtg Val	cgg Arg	gtg Val	tcg Ser 45	gcc Ala	gag Glu	gtc Val	gct Ala	tac Tyr 50	cgc Arg	tcg Ser	498
ccg Pro	acg Thr	ggc Gly 55	gcc Ala	cac His	Gly 999	ttc Phe	ggc Gly 60	ccg Pro	atc	cgc Arg	gtc Val	ctc Leu 65	gat Asp	gcc Ala	gag Glu	546
ggt Gly	gtg Val 70	ccg Pro	gtg Val	gcg Ala	ctc Leu	gcc Ala 75	gat Asp	ccg Pro	gtg Val	ctg Leu	ctg Leu 80	gcg Ala	gcc Ala	gcc Ala	tgc Cys	594
tcg Ser 85	Ala	gac Asp	tcg Ser	cgg Arg	agc Ser 90	cgc Arg	tcg Ser	ctg Leu	ccg Pro	agc Ser 95	Ala	ccg Pro	atc Ile	aac Asn	gcc Ala 100	642
ccg Pro	gac Asp	gcc Ala	ggt Gly	acc Thr 105	gct Ala	gtc Val	gac Asp	tgg Trp	gtg Val 110	Leu	tcg Ser	tcg Ser	ctc Leu	gcc Ala 115	gac Asp	690
gac Asp	gag Glu	gac Asp	gac Asp 120	Glu	gtg Val	ccc Pro	gcc Ala	ggc Gly 125	Met	acc Thr	gcg Ala	gag Glu	gag Glu 130	gcg Ala	gtg Val	738
cgc Arc	ctg Leu	ctg Leu 135	Ser	cgg Arg	cag Gln	gtc Val	gac Asp 140	Asp	ctg Leu	ccg Pro	g cgg Arg	tcg Ser 145	PIC	ggc Gly	gcc Ala	786
gad Asp	c ccg Pro 150	Trp	tcg Ser	g ctg Leu	gto Val	gcc Ala 155	Gly	e ccg	g cto Lei	g gcg n Ala	g gco a Ala 160	я тте	: Gl <sup>)</sup>	g cgg y Arg	ttc Phe	834
999 Gl <sub>3</sub> 169	/ Arg	gcc Ala	ggg Gl	g ato / Ile	gcc Ala 170	ı Asp	gag Glu	g tgo i Cys	e tgg s Tri	tto Lev 17	л ге	g gag ı Glu	g gtg i Val	g cto l Lev	gcc Ala 180	882



GJÀ 333	cgg Arg	ctc Leu	cgc Arg	gcg Ala 185	gtc Val	gac Asp	gac Asp	gac Asp	ctg Leu 190	tcc Ser	cgc Arg	tcg Ser	tgg Trp	ctg Leu 195	ago Ser	: :	930
agt Ser	ccg Pro	acg Thr	ctc Leu 200	gcc Ala	gac Asp	cgc Arg	gct Ala	gtg Val 205	ctc Leu	gtg Val	ggt Gly	gag Glu	999 Gly 210	ttg Leu	Arg	3	978
tac Tyr	cgg Arg	ccg Pro 215	gat Asp	gtg Val	cgg Arg	ccg Pro	gtg Val 220	ccg Pro	ttc Phe	gac Asp	gtg Val	ccg Pro 225	aac Asn	ccg Pro	Le	g u	1026
cac His	gag Glu 230	ggc	aag Lys	tcc Ser	gac Asp	gtc Val 235	Pro	ccg Pro	ccg Pro	ccc Pro	gtg Val 240	FIU	gtg Val	ctg Leu	gg Gl	с У	1074
999 Gly 245	ccg Pro	tgg Trp	tcg Ser	ctg Leu	cgt Arg 250	Pro	gtc Val	gag Glu	gtc Val	gcg Ala 255	· vai	cac His	Gly Ggg	gat Asp	gg Gl 26	4	1122
gly aaa	cct Pro	gac Asp	gto Val	gca Ala 265	Leu	gtg Val	cac His	cgc	tgg Trp 270	Mec	aac Asn	acc Thr	ccg Pro	cac His		c al	1170
gcg Ala	cac His	Cac	tgg Trp 280	g aac o Asn	cag Glr	gcg Ala	tgg Trp	ccg Pro 285	, пес	gag ıGlı	g cgo	tgg Tr	cgg Arg 290	,	g ga	aa lu	1218
cto	gcc Ala	cac His	s Gli	g cto n Lev	ggo Gly	ggt Gly	gag Glu 300	1 HIS	tco Sei	c cto	ı Pro	c tgo c Cy:	. vu	g gte l Va	c gg	ga ly	1266
cac His	gaç Glu	ı Gl	a cg y Ar	c gaq g Gli	g gto ı Va	gc l Ala 31	а ту:	t cto	g gaq ı Glı	g cto u Le	c ta u Ty 32	_ A.	g gte g Va	g ac l Th	c c r A	gc rg	1314
gae Asj 32!	aaq b Lys	ndII g ct s Le	- ~~	g gg a Gl	c tg y Cy 33	в Ту	c cc r Pr	g ta o Ty:	c gg r Gl	g cc y Pr 33	O AI	c ga s As	c ct p Le	c gg u Gl	1 .	tc al 40	1362
ca Hi	c ato	c gc e Al	g at a Il	c gg e Gl 34	λ GT	g cg u Ar	g ga g Gl	g gt u Va	g ct 1 Le 35	u Gi	g cg y Ar	g Gl	t tt y Ph	c gg le Gl 35	. 4 ~	cg	1410
tc Se	g ct r Le	g ct u Le	g cg u Ar 36	g Al	g gt a Va	c gc	g gg .a Gl	t go y Al	а ье	g ct	g ga eu As	c go	c ga a As	יב עי	eg o	egg Arg	1458
tg Cy	c gc s Al	g cg a Ai 37	g Va	g gt al Va	c go	c ga .a Gl	ig co lu Pi 38	O AS	it gt in Va	g ca al Hi	ac aa is As	311 0.	ag go Lu Al 35	ct to la Se	eg 9	gtg Val	1506



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cgc gcc ttc gcc aag Arg Ala Phe Ala Lys 390	gcc ggg ttc g Ala Gly Phe N 395	gtc cgg gag Val Arg Glu	agg gag atc gg Arg Glu Ile Gl 400	c ctg 1554 y Leu
ccc gcc aag aac tcg Pro Ala Lys Asn Ser 405	gct ctg atg of Ala Leu Met 1410	gtc ttc tcc Val Phe Ser 415	cgg gtc t gacg Arg Val (SEQ I	accggt 1604 D NO:2)
catgccctg tgtgaacgc ggttagcctt acttttatt ctgaccacag cgcagtagg tacaacaccg gcgagtgcg gccctgcggt ggggcgtgg gacgagccgc gccaggtcg gcgccgaggc ggaccc (S	g gtggagaacg g catctggttc gt cgagatcagc gc ggaggccgag gt ggagccgag	atgccggagc gcccagcaac ggcccggtgg gcgctgcgag	gctccgccgt gtc tcgaccggac gaa agccggtggt gtt cccgcgtggt cgt	gttgccg 1724 cccgatc 1784 cgagcag 1844 cgacggc 1904



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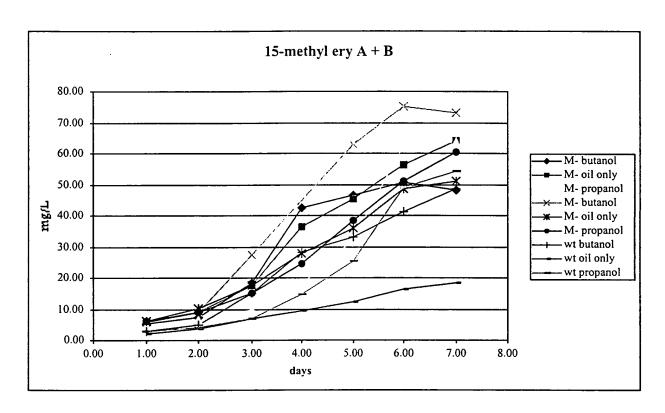
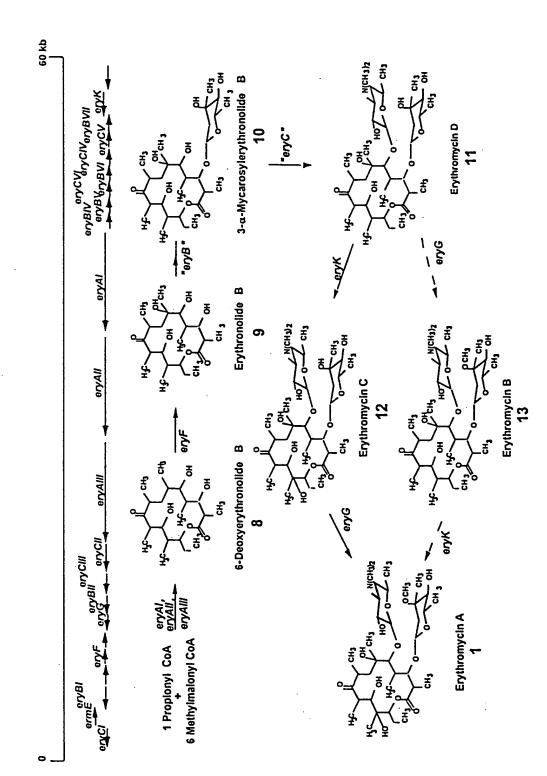


Figure 4



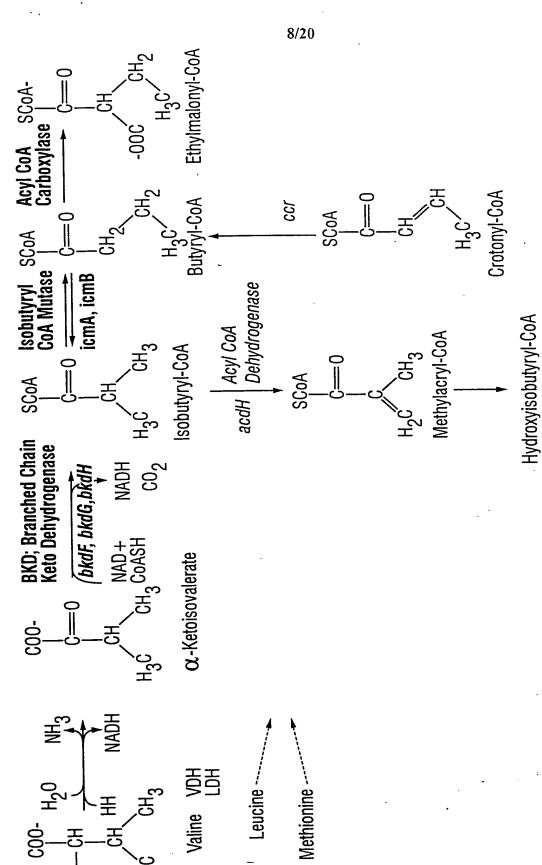


Figure 6 A



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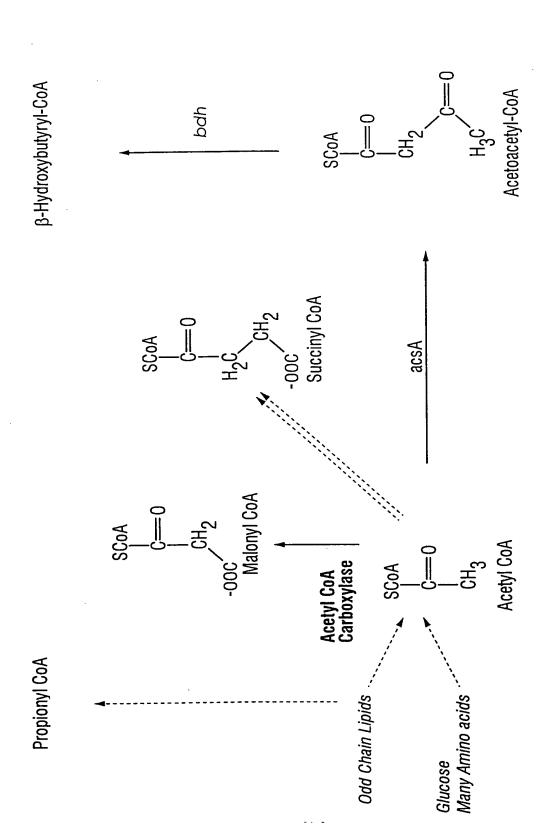


Figure 6 C



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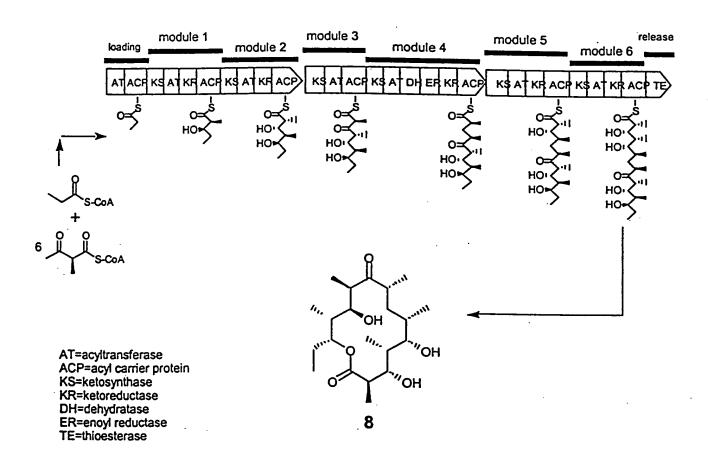


Figure 7



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Figure 8



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Figure 9



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Met His Val Pro Gly Glu Glu Asn Gly His Ser Ile Ala Ile Val Gly 10 Ile Ala Cys Arg Leu Pro Gly Ser Ala Thr Pro Gln Glu Phe Trp Arg 25 Leu Leu Ala Asp Ser Ala Asp Ala Leu Asp Glu Pro Pro Ala Gly Arg 40 Phe Pro Thr Gly Ser Leu Ser Ser Pro Pro Ala Pro Arg Gly Gly Phe Leu Asp Ser Ile Asp Thr Phe Asp Ala Asp Phe Phe Asn Ile Ser Pro Arg Glu Ala Gly Val Leu Asp Pro Gln Gln Arg Leu Ala Leu Glu Leu 90 Gly Trp Glu Ala Leu Glu Asp Ala Gly Ile Val Pro Arg His Leu Arg 105 100 Gly Thr Arg Thr Ser Val Phe Met Gly Ala Met Trp Asp Asp Tyr Ala 120 His Leu Ala His Ala Arg Gly Glu Ala Ala Leu Thr Arg His Ser Leu 140 135 Thr Gly Thr His Arg Gly Met Ile Ala Asn Arg Leu Ser Tyr Ala Leu 155 150 Gly Leu Gln Gly Pro Ser Leu Thr Val Asp Thr Gly Gln Ser Ser 170 Leu Ala Ala Val His Met Ala Cys Glu Ser Leu Ala Arg Gly Glu Ser 185 Asp Leu Ala Leu Val Gly Gly Val Asn Leu Val Leu Asp Pro Ala Gly 200 Thr Thr Gly Val Glu Arg Phe Gly Ala Leu Ser Pro Asp Gly Arg Cys 215 Tyr Thr Phe Asp Ser Arg Ala Asn Gly Tyr Ala Arg Gly Glu Gly Gly 235 230 Val Val Val Leu Lys Pro Thr His Arg Ala Leu Ala Asp Gly Asp 250 245 Thr Val Tyr Cys Glu Ile Leu Gly Ser Ala Leu Asn Asn Asp Gly Ala 265 Thr Glu Gly Leu Thr Val Pro Ser Ala Arg Ala Gln Ala Asp Val Leu 280 Arg Gln Ala Trp Glu Arg Ala Arg Val Ala Pro Thr Asp Val Gln Tyr 300 295 Val Glu Leu His Gly Thr Gly Thr Pro Ala Gly Asp Pro Val Glu Ala 315 310 Glu Gly Leu Gly Thr Ala Leu Gly Thr Ala Arg Pro Ala Glu Ala Pro 330 Leu Leu Val Gly Ser Val Lys Thr Asn Ile Gly His Leu Glu Gly Ala 345 Ala Gly Ile Ala Gly Leu Leu Lys Thr Val Leu Ser Ile Lys Asn Arg 360 His Leu Pro Ala Ser Leu Asn Phe Thr Ser Pro Asn Pro Arg Ile Asp 380 375 Leu Asp Ala Leu Arg Leu Arg Val His Thr Ala Tyr Gly Pro Trp Pro 395 390 Ser Pro Asp Arg Pro Leu Val Ala Gly Val Ser Ser Phe Gly Met Gly 405 410



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Gly	Thr	Asn	Cys 420	His	Val	Val	Leu	Ser 425	Glu	Leu	Arg	Asn	Ala 430	Gly	Gly
Asp	Gly	Ala 435		Lys	Gly	Pro	Tyr 440		Gly	Thr	Glu	Asp 445	Arg	Leu	Gly
	450	Glu			Lys	455					460				
465					Thr 470					475					480
_				485	Leu				490					495	
			500		Gly			505					510		
		515			Val		520					525			
_	530				Leu	535					540				
545					Leu 550					555					560
_				565	Phe				570					575	
_			580		Ala			585					590		
	_	595			Leu		600					605			
	610	•			Glu	615					620				
625	-	_			Met 630					635					640
_				645	Ala				650					655	•
			660		Gly			665					670		
		675			Val		680					685			
_	690				Val	695					700				
705		_			710					715					Thr 720
				725	Gly				730					735	
			740					745					750		Cys
		755					760					765			Arg
	770		_			775					780				Ser
785					790					795					Trp 800
				805					810					815	
Ala	Arg	Glu	Pro 820		Asp	Thr	val	Phe 825	val	GIU	val	ser	830		Pro



Val	Leu	Leu 835	Pro	Ala	Ile	Asp	Gly 840	Ala	Thr	Val	Ala	Thr 845	Leu	Arg	Arg
Gly	Gly 850	Gly	Val	His	Arg	Leu 855	Leu	Thr	Ala	Leu	Ala 860	Glu	Ala	His	Thr
Thr 865	Gly	Val	Pro	Val	Asp 870	Trp	Ala	Ala	Val	Val 875	Pro	Ala	Thr	Ala	Thr 880
Ala	His	Asp	Leu	Pro 885	Thr	Tyr	Ala	Phe	His 890	His	Glu	Arg	Tyr	Trp 895	Ile
Ser	His	Trp	Leu 900	Pro	Ser	Gly	Glu	Ala 905	His	Pro	Arg	Pro	Ala 910	Asp	Asp
Thr	Glu	Ser 915	Gly	Thr	Gly	Arg	Thr 920	Glu	Ala	Ser	Pro	Pro 925	Arg	Pro	His
Asp	(SE	Q ID	NO:	3)											



			_				_			_	_				
Met 1	His	Val	Pro	Gly 5	Glu	Glu	Asn	GIY	GIu 10	Pro	Leu	Ala	Ile	Val 15	Gly
Met	Ala	Cys	Arg 20	Leu	Pro	Gly	Gly	Val 25	Ala	Ser	Pro	Glu	Asp 30	Leu	Trp
Arg	Leu	Leu 35	Glu	Ser	Gly	Gly	Asp 40	Gly	Ile	Thr	Ala	Phe 45	Pro	Thr	Asp
Arg	Gly 50	Trp	Asp	Val	Asp	Gly 55	Leu	Tyr	Asp	Pro	Asp 60	Pro	Asp	His	Pro
Gly 65		Ser	Thr	Val	Arg 70		Gly	Gly	Phe	Leu 75	Ala	Gly	Val	Ala	Asp 80
	Asp	Ala	Ala	Phe 85		Gly	Ile	Ser	Pro 90	Arg	Glu	Ala	Leu	Ala 95	
Asp	Pro	Gln	Gln 100		Leu	Val	Leu	Glu 105	Thr	Ser	Trp	Glu	Ala 110		Glu
His	Ala	Gly 115	Ile	Leu	Pro	Glu	Ser 120		Arg	Gly	Ser	Asp 125		Gly	Val
Phe	Met 130		Ala	Phe	Ser	Asp		Tyr	Gly	Leu	Gly 140		Asp	Leu	Gly
Gly 145		Gly	Ala	Thr	Gly 150		Gln	Thr	Ser	Val 155		Ser	Gly	Arg	Leu 160
	Tyr	Phe	Tyr	Gly 165		Glu	Gly	Pro	Ala 170		Thr	Val	Asp	Thr 175	
Cys	Ser	Ser	Ser 180	Leu	Val	Ala	Leu	His 185	Gln	Ala	Gly	Gln	Ser 190	Leu	Arg
Ser	Gly	Glu 195	Cys	Ser	Leu	Ala	Leu 200	Val	Gly	Gly	Val	Thr 205	Val	Met	Ala
Ser	Pro 210	Ser	Gly	Phe	Val	Glu 215	Phe	Ser	Gln	Gln	Arg 220	Gly	Leu	Ala	Pro
Asp 225	Äla	Arg	Cys	Lys	Ala 230	Phe	Ala	Asp	Ala	Ala 235	Asp	Gly	Thr	Gly	Phe 240
Ala	Glu	Gly	Ser	Gly 245	Val	Leu	Ile	Val	Glu 250	Arg	Leu	Ser	Asp	Ala 255	Glu
Arg	Asn	Gly	His 260	Arg	Val	Leu	Ala	Val 265	Val	Arg	Gly	Ser	Ala 270	Val	Asn
	_	275	Ala		Asn	Gly		Ser	Ala	Pro	Asn		Pro	Ser	Gln
Glu	_	1/2 l					280					285			
	290	Val	Ile	Arg	Gln	Ala 295		Ala	Asn	Ala	Gly 300		Thr	Pro	Ala
Asp 305	Val		Ala	Val		295 Ala	Leu His		Thr	Gly	300 Thr	Leu			
305 Pro	Val Ile	Asp Glu	Ala Ala	Val Gln 325	Glu 310 Ala	295 Ala Val	Leu His Leu	Gly Ala	Thr Thr 330	Gly 315 Tyr	300 Thr Gly	Leu Arg Gln	Leu Gly	Gly Arg 335	Asp 320 Asp
305 Pro Thr	Val Ile Pro	Asp Glu Val	Ala Ala Leu 340	Val Gln 325 Leu	Glu 310 Ala Gly	295 Ala Val Ser	Leu His Leu Leu	Gly Ala Lys 345	Thr Thr 330 Ser	Gly 315 Tyr Asn	300 Thr Gly Ile	Leu Arg Gln Gly	Leu Gly His 350	Gly Arg 335 Thr	Asp 320 Asp Gln
305 Pro Thr	Val Ile Pro Ala	Asp Glu Val Ala 355	Ala Ala Leu 340 Gly	Val Gln 325 Leu Val	Glu 310 Ala Gly Ala	295 Ala Val Ser Gly	Leu His Leu Leu Val 360	Gly Ala Lys 345 Ile	Thr Thr 330 Ser Lys	Gly 315 Tyr Asn Met	300 Thr Gly Ile Val	Leu Arg Gln Gly Leu 365	Leu Gly His 350 Ala	Gly Arg 335 Thr	Asp 320 Asp Gln Arg
305 Pro Thr Ala His	Val Ile Pro Ala Gly 370	Asp Glu Val Ala 355 Thr	Ala Ala Leu 340 Gly Leu	Val Gln 325 Leu Val Pro	Glu 310 Ala Gly Ala Arg	295 Ala Val Ser Gly Thr 375	Leu His Leu Leu Val 360 Leu	Gly Ala Lys 345 Ile	Thr Thr 330 Ser Lys Val	Gly 315 Tyr Asn Met	300 Thr Gly Ile Val Thr 380	Leu Arg Gln Gly Leu 365 Pro	Leu Gly His 350 Ala Ser	Gly Arg 335 Thr Met Ser	Asp 320 Asp Gln Arg
305 Pro Thr Ala His Val 385	Val Ile Pro Ala Gly 370 Asp	Asp Glu Val Ala 355 Thr	Ala Leu 340 Gly Leu Thr	Val Gln 325 Leu Val Pro	Glu 310 Ala Gly Ala Arg Gly 390	295 Ala Val Ser Gly Thr 375 Ala	Leu His Leu Leu Val 360 Leu Val	Gly Ala Lys 345 Ile His Glu	Thr Thr 330 Ser Lys Val Leu	Gly 315 Tyr Asn Met Asp Leu 395	300 Thr Gly Ile Val Thr 380 Thr	Leu Arg Gln Gly Leu 365 Pro	Leu Gly His 350 Ala Ser	Gly Arg 335 Thr Met Ser Arg	Asp 320 Asp Gln Arg His
305 Pro Thr Ala His Val 385 Trp	Val Ile Pro Ala Gly 370 Asp	Asp Glu Val Ala 355 Thr Trp	Ala Leu 340 Gly Leu Thr	Val Gln 325 Leu Val Pro Ala Asp 405	Glu 310 Ala Gly Ala Arg Gly 390 Arg	295 Ala Val Ser Gly Thr 375 Ala Pro	Leu His Leu Val 360 Leu Val	Gly Ala Lys 345 Ile His Glu Arg	Thr 330 ser Lys Val Leu Ala 410	Gly 315 Tyr Asn Met Asp Leu 395 Gly	300 Thr Gly Ile Val Thr 380 Thr	Leu Arg Gln Gly Leu 365 Pro Asp Ser	Leu Gly His 350 Ala Ser Ala Ser	Gly Arg 335 Thr Met Ser Arg Phe 415	Asp 320 Asp Gln Arg His Pro 400 Gly
305 Pro Thr Ala His Val 385 Trp	Val Ile Pro Ala Gly 370 Asp Pro	Asp Glu Val Ala 355 Thr Trp Glu Gly	Ala Leu 340 Gly Leu Thr Thr	Val Gln 325 Leu Val Pro Ala Asp 405 Asn	Glu 310 Ala Gly Ala Arg Gly 390 Arg	295 Ala Val Ser Gly Thr 375 Ala Pro	Leu His Leu Val 360 Leu Val Arg	Gly Ala Lys 345 Ile His Glu Arg Leu 425	Thr 330 Ser Lys Val Leu Ala 410 Leu	Gly 315 Tyr Asn Met Asp Leu 395 Gly	300 Thr Gly Ile Val Thr 380 Thr Val	Leu Arg Gln Gly Leu 365 Pro Asp Ser His	Leu Gly His 350 Ala Ser Ala Ser Pro 430	Gly Arg 335 Thr Met Ser Arg Phe 415 Ala	Asp 320 Asp Gln Arg His Pro 400 Gly
305 Pro Thr Ala His Val 385 Trp	Val Ile Pro Ala Gly 370 Asp Pro	Asp Glu Val Ala 355 Thr Trp Glu Gly	Ala Leu 340 Gly Leu Thr Thr	Val Gln 325 Leu Val Pro Ala Asp 405 Asn	Glu 310 Ala Gly Ala Arg Gly 390 Arg	295 Ala Val Ser Gly Thr 375 Ala Pro	Leu His Leu Val 360 Leu Val Arg	Gly Ala Lys 345 Ile His Glu Arg Leu 425	Thr 330 Ser Lys Val Leu Ala 410 Leu	Gly 315 Tyr Asn Met Asp Leu 395 Gly	300 Thr Gly Ile Val Thr 380 Thr Val	Leu Arg Gln Gly Leu 365 Pro Asp Ser His	Leu Gly His 350 Ala Ser Ala Ser Pro 430	Gly Arg 335 Thr Met Ser Arg Phe 415 Ala	Asp 320 Asp Gln Arg His Pro 400 Gly



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Ile	Ala 450	Thr	Pro	Leu	Thr	Pro 455	Leu	Pro	Val	Ser	Ala 460	Arg	Thr	Ala	Thr
Ala 465	Leu	Asp	Gly	Gln	Val 470		Arg	Leu	Arg	Glu 475	His	Leu	Ala	Ala	Arg 480
	Gly	His	Asp	Pro		Ala	Ile	Ala	Ala 490		Leu	Leu	Ala	Arg 495	
Thr	Thr	Phe			Arg	Ala	Val	Leu 505		Asp	Asp	Asp	Val 510		Thr
Gly	Thr		500 Leu	Thr	Glu	Pro			Val	Phe	Val			Gly	Gln
Gly	Pro	515 Gln	Trp	Arg	Gly		520 Gly	Val	Glu	Leu		525 Ala	Ala	Ser	Pro
	530					535	_		_		540		_		
Val	Phe	Ala	Ala	Arg		Arg	Gln	Cys	Ala		Ala	Leu	Ile	Pro	
545					550					555					560
	Gly	_	-	565					570					575	
_	Val	_	580					585					590		
Ala	Ala	Val	Trp	Glu	Ala	Ala	Gly	Val	Arg	Pro	Asp	Ala	Val	Ile	Gly
		595					600					605			
His	Ser	Gln	Gly	Glu	Ile	Ala	Ala	Ala	Cys	Val	Ala	Gly	Ala	Leu	Thr
	610		_			615					620				
Leu	Glu	Asp	Glv	Ala	Arq	Leu	Val	Ala	Leu	Arg	Ser	Val	Leu	Leu	Leu
625			1		630					635					640
	Arg	Glu	Len	Δla		Ara	Glv	Ala	Met		Ser	Val	Ala	Leu	
шeu	Arg	Giu	пец	645	Gry	T. A	Ory	AIG	650		001	<b>V</b> 41	7114	655	110
77-	Ala	7 ~~	1707		. ד מ	λαn	- ד ג	πΙэ		Tla	Acn	Glv	V-1		17 <b>-</b> 1
мта	Ата	Asp		GIU	AIA	Asp	AIG	665	Arg	110	Hop	Cry	670	ııp	var
ח ח	Gly	7	660	a1	71-	Th.~	Thr		Thr	V-1	λla	Glaz		Dro	Λen
	_	675					680					685			
	Val 690					695					700				
	Arg	Ile	Ala	Val		Cys	Pro	Thr	His		Pro	Phe	Val	Asp	
705					710	_				715	_,	_,	_	_	720
	Tyr	_		725					730					735	
	Glu		740					745					750		
Pro	Leu	Asp 755	_	Glu	Tyr		Phe 760		Asn			His 765		Val	Gly
Phe	Ala 770	Thr	Ala	Val	Thr	Ala 775	Ala	Arg	Glu	Pro	Gly 780	Asp	Thr	Val	Phe
Val 785	Glu	Val	Ser	Ala	His 790	Pro	Val	Leu	Leu	Pro 795	Ala	Ile	Asp	Gly	Ala 800
	Val	Ala	Thr	Leu		Ara	Glv	Glv	Glv	Val	His	Arq	Leu	Leu	Thr
				805	5	3	1	4	810					815	
λ] -	Leu	λla	Glu		Hie	Thr	Thr	Glv			Val	Asp	Tro		Δla
AIO	пец	AIG	820	AIG	1113	1111		825		110			830		1114
17- 1	**- 1	D		mb	77-		<b>7.7</b> -			Lou	Dro	Thr		ת ד ת	Dhe
val	Val		МТЯ	TIIL	HIG	TIIL		1112	чэb	neu	210	845	+ A T	ALA	r 116
		835		<b></b>		<b>~</b> 7.	840	77.2 -	П	T	D		<b>01.</b>	<b>71.</b>	7.7.
His	His	Glu	Arg	Tyr	Trp		ser	HlS	Trp	ьeu		ser	GTĀ	GIU	АТА
	850		_		_	855		~ 7	~	<b>~</b> 3	860	a 3		m'	a 3
	Pro	Arg	Pro	Ala			Thr	Glu	Ser		Thr	GIY	Arg	Thr	
865					870		_			875					880
Ala	Ser	Pro	Pro	Arg 885	Pro	His	Asp	(SE	Q ID	NO:	4)				



Inventor: Leonard KATZ and Peter REVILL Title: PRODUCTION OF POLYKETIDES

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Met His Val Pro Gly Glu Glu Asn Gly Glu Pro Leu Ala Ile Val Gly 1.0 Met Ala Cys Arg Leu Pro Gly Gly Val Ala Ser Pro Glu Asp Leu Trp Arg Leu Leu Glu Ser Gly Gly Asp Gly Ile Thr Ala Phe Pro Thr Asp 40 Arg Gly Trp Asp Val Asp Gly Leu Tyr Asp Pro Asp Pro Asp His Pro 60 Gly Thr Ser Thr Val Arg His Gly Gly Phe Leu Ala Gly Val Ala Asp 75 Phe Asp Ala Ala Phe Phe Gly Ile Ser Pro Arg Glu Ala Leu Ala Met Asp Pro Gln Gln Arg Leu Val Leu Glu Thr Ser Trp Glu Ala Leu Glu 105 His Ala Gly Ile Leu Pro Glu Ser Leu Arg Gly Ser Asp Thr Gly Val 120 125 Phe Met Gly Ala Phe Ser Asp Gly Tyr Gly Leu Gly Thr Asp Leu Gly 140 135 Gly Phe Gly Ala Thr Gly Thr Gln Thr Ser Val Leu Ser Gly Arg Leu 155 150 Ser Tyr Phe Tyr Gly Leu Glu Gly Pro Ala Val Thr Val Asp Thr Ala 165 170 Gln Ser Ser Leu Val Ala Leu His Gln Ala Gly Gln Ser Leu Arg 185 Ser Gly Glu Cys Ser Leu Ala Leu Val Gly Gly Val Thr Val Met Ala 200 Ser Pro Ser Gly Phe Val Glu Phe Ser Gln Gln Arg Gly Leu Ala Pro 215 Asp Ala Arg Cys Lys Ala Phe Ala Asp Ala Ala Asp Gly Thr Gly Phe 235 230 Ala Glu Gly Ser Gly Val Leu Ile Val Glu Arg Leu Ser Asp Ala Glu 250 Arg Asn Gly His Arg Val Leu Ala Val Val Arg Gly Ser Ala Val Asn 265 Gln Asp Gly Ala Ser Asn Gly Leu Ser Ala Pro Asn Gly Pro Ser Gln 280 Glu Arg Val Ile Arg Gln Ala Leu Ala Asn Ala Gly Leu Thr Pro Ala 300 295 Asp Val Asp Ala Val Glu Ala His Gly Thr Gly Thr Arg Leu Gly Asp 310 315 Pro Ile Glu Ala Gln Ala Val Leu Ala Thr Tyr Gly Gln Gly Arg Asp 330 325 Thr Pro Val Leu Leu Gly Ser Leu Lys Ser Asn Ile Gly His Thr Gln 345 Ala Ala Gly Val Ala Gly Val Ile Lys Met Val Leu Ala Met Arg 360 His Gly Thr Leu Pro Arg Thr Leu His Val Asp Thr Pro Ser Ser His 375 380 Val Asp Trp Thr Ala Gly Ala Val Glu Leu Leu Thr Asp Ala Arg Pro 395 Trp Pro Glu Thr Asp Arg Pro Arg Arg Ala Gly Val Ser Ser Phe Gly 410 405 Val Ser Gly Thr Asn Ala His Val Leu Leu Glu Ala His Pro Ala Gly 425 Glu Pro Pro Ala Glu Glu Pro Ser Ala Ser Lys Pro Gly Glu Pro Leu 440 445 435



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Ile Ala Thr Pro Leu Thr Pro Leu Pro Val Ser Ala Arg Thr Ala Thr 455 Ala Leu Asp Gly Gln Val Arg Arg Leu Arg Glu His Leu Ala Ala Arg 475 470 Pro Gly His Asp Pro Arg Ala Ile Ala Ala Gly Leu Leu Ala Arg Arg 485 490 Thr Thr Phe Pro His Arg Ala Val Leu Leu Asp Asp Val Val Thr 505 500 Gly Thr Ala Leu Thr Glu Pro Arg Thr Val Phe Val Phe Pro Gly Gln 520 Gly Pro Gln Trp Arg Gly Met Gly Val Glu Leu Met Ala Ala Ser Pro 535 Val Phe Ala Ala Arg Met Arg Gln Cys Ala Asp Ala Leu Ile Pro His 555 550 Thr Gly Trp Asp Pro Ile Ala Met Leu Asp Asp Pro Glu Val Thr Arg 570 565 Arg Val Asp Val Val His Pro Val Cys Trp Ala Val Met Val Ser Leu 585 Ala Ala Val Trp Glu Ala Ala Gly Val Arg Pro Asp Ala Val Ile Gly 600 His Ser Gln Gly Glu Ile Ala Ala Cys Val Ala Gly Ala Leu Thr 615 Leu Glu Asp Gly Ala Arg Leu Val Ala Leu Arg Ser Val Leu Leu 630 635 Leu Arg Glu Leu Ala Gly Arg Gly Ala Met Gly Ser Val Ala Leu Pro 645 650 Ala Ala Asp Val Glu Ala Asp Ala Ala Arg Ile Asp Gly Val Trp Val Ala Gly Arg Asn Gly Ala Thr Thr Thr Thr Val Ala Gly Arg Pro Asp 680 Ala Val Glu Thr Leu Ile Ala Asp Tyr Glu Ala Arg Gly Val Trp Val 695 Arg Arg Ile Ala Val Asp Cys Pro Thr His Thr Pro Phe Val Asp Pro 715 710 Leu Tyr Asp Glu Leu Gln Arg Ile Val Ala Asp Thr Thr Ser Arg Thr 730 725 Pro Glu Ile Pro Trp Phe Ser Thr Ala Asp Glu Arg Trp Ile Asp Ala 745 740 Pro Leu Asp Asp Glu Tyr Trp Phe Arg Asn Met Arg His Pro Val Gly 760 Phe Ala Thr Ala Val Thr Ala Ala Arg Glu Pro Gly Asp Thr Val Phe 780 775 Val Glu Val Ser Ala His Pro Val Leu Leu Pro Ala Ile Asp Gly Ala 790 795 Thr Val Ala Thr Leu Arg Arg Gly Gly Val His Arg Leu Leu Thr 805 810 Ala Leu Ala Glu Ala His Thr Thr Gly Val Pro Val Asp Trp Ala Ala 825 820 Val Val Pro Ala Thr Ala Thr Ala His Asp Leu Pro Thr Tyr Ala Phe 840 845 His His Glu Arg Tyr Trp Ile Ser His Trp Leu Pro Ser Gly Glu Ala 855 860 His Pro Arg Pro Ala Asp Asp Thr Glu Ser Gly Thr Gly Arg Thr Glu 870 Ala Ser Pro Pro Arg Pro His Asp (SEQ ID NO:5) 885